

**EXPRESS MAIL LABEL NO. EL 979108920 US
ATTORNEY DOCKET NO. 07082.0016U1
Application No. PCT/AU2004/000474**

Listing of Claims

Claims 1-40 (amended under Article 34) (canceled).

41. (New) A method of treatment or prophylaxis of a condition associated with elevated levels of non-amidated gastrin, comprising the step of administering to a mammal in need of such treatment an effective amount of a compound which has the ability to inhibit the binding of ferric ions to any one or more of glycine-extended gastrin₁₇ or progastrin or progastrin-derived peptides, but which does not inhibit the activity of amidated gastrin, thereby to inhibit the activity of non-amidated gastrins.

42. (New) A method according to claim 41, in which the compound inhibits the binding of ferric ions to glutamate 7 of glycine-extended gastrin₁₇.

43. (New) A method according to claim 42, in which the binding of ferric ions to glutamate 8 and glutamate 9 of glycine-extended gastrin₁₇ is also inhibited.

44. (New) A method according to claim 41, in which the compound is a metal ion, or a pharmaceutically-acceptable salt or complex thereof, which is able to occupy the ferric ion binding site of non-amidated gastrins, and thereby to block their biological activity.

45. (New) A method according to claim 44, in which the metal ion is any metal ion capable of occupying the ferric ion binding site of non-amidated gastrins, with the provisos that

(i) when the condition is one caused by *Helicobacter pylori* infection, the metal ion is not bismuth, and

(ii) when the condition is cancer, the salt or complex is not BiSrC₆H₅O₆.

46. (New) A method according to claim 45, in which the metal ion is Bi³⁺ or Ga³⁺.

**EXPRESS MAIL LABEL NO. EL 979108920 US
ATTORNEY DOCKET NO. 07082.0016U1
Application No. PCT/AU2004/000474**

47. (New) A method according to claim 41, in which the compound is an exchange-inert complex between a non-amidated gastrin and either Co (III) or Cr (III) ions.

48. (New) A method according to claim 41, in which the compound is a pharmaceutically-acceptable chelating agent with a high degree of specificity for ferric ions.

49. (New) A method according to claim 48, in which the chelating agent is membrane-impermeable.

50. (New) A method according to claim 49, in which the chelating agent is desferrioxamine (DFO), ethylene diamine tetracetic acid (EDTA) or diethylene triamine pentacetic acid (DTPA).

51. (New) A method according to claim 48, in which the chelating agent is a membrane-permeable chelator.

52. (New) A method according to claim 51, in which the chelating agent is clioquinol.

53. (New) A method according to claim 41, in which the compound does not have a significant inhibitory effect on Gamide-induced inositol phosphate production and/or on cellular proliferation in cells which express the CCK-2 receptor.

54. (New) A method according to claim 46, in which the compound is one or more of colloidal bismuth subcitrate (CBS), bismuth subcitrate, bismuth citrate, bismuth salicylate, bismuth subsalicylate, bismuth subnitrate, bismuth subcarbonate, bismuth tartrate, bismuth subgallate, tripotassium dicitrato bismuthate or bismuth aluminate.

**EXPRESS MAIL LABEL NO. EL 979108920 US
ATTORNEY DOCKET NO. 07082.0016U1
Application No. PCT/AU2004/000474**

55. (New) A method according to claim 54, in which the compound is one or more of colloidal bismuth subcitrate (CBS), tripotassium dicitrato bismuthate, bismuth subcitrate, or bismuth subsalicylate.

56. (New) A method according to claim 55, in which the compound is CBS or tripotassium dicitrato bismuthate.

57. (New) A method according to claim 56, in which the compound is CBS.

58. (New) A method according to claim 41, in which the condition is selected from the group consisting of gastrin-producing tumours, colorectal carcinomas, gastrinomas, islet cell carcinomas, lung cancer, ovarian cancer, pituitary cancer and pancreatic cancer.

59. (New) A method according to claim 58, in which the condition is colon cancer or pancreatic cancer.

60. (New) A method according to claim 59, in which the condition is colon cancer and the mammal is at elevated risk thereof.

61. (New) A method according to claim 60, in which the mammal is an individual with any one or more of familial adenomatous polyposis, with a family history of colon cancer, and/or with loss of imprinting of IGF-2.

62. (New) A method according to claim 41, in which the condition is selected from the group consisting of atrophic gastritis, G cell hyperplasia, pernicious anaemia, renal failure and ulcerative colitis.

**EXPRESS MAIL LABEL NO. EL 979108920 US
ATTORNEY DOCKET NO. 07082.0016U1
Application No. PCT/AU2004/000474**

63. (New) A method according to claim 41, in which the condition is selected from the group consisting of gastrointestinal ulcers, gastro-oesophageal reflux, gastric carcinoid, and Zollinger-Ellison syndrome, with the proviso that the metal ion is not bismuth.

64. (New) A peptide which is a fragment of a non-amidated gastrin and which

(a) comprises at least glutamate residue 7 of the -(Glu)₅- sequence of non-amidated gastrin, and

(b) is capable of binding one or more ferric ions, with the proviso that the peptide is not full length Ggly, full length glycine-extended gastrin or full length progastrin, or LE₅AYG.

65. (New) A peptide according to claim 64, consisting of amino acids 5 to 14 of the Ggly sequence.

66. (New) A peptide according to claim 64, selected from the group consisting of Ggly₅₋₁₈, Ggly₁₋₁₁, LE5AY, LE₅A, LE₅E, E₅A, and E₅AY.

67. (New) A peptide according to claim 64, in which the carboxy terminus of the peptide is amidated.

68. (New) A peptide according to claim 64, in which the amino terminus of the peptide is acetylated.

69. (New) A complex comprising

(a) a non-amidated gastrin, a peptide fragment thereof according to claim 64, or LE₅AYG, and

(b) a trivalent metal ion.

**EXPRESS MAIL LABEL NO. EL 979108920 US
ATTORNEY DOCKET NO. 07082.0016U1
Application No. PCT/AU2004/000474**

70. (New) A complex according to claim 69, in which the trivalent metal ion is Bi^{3+} or Ga^{3+} .

71. (New) A complex according to claim 69, comprising a non-amidated gastrin and bismuth ions.

72. (New) A composition comprising

(a) a peptide according to claim 64, or LE₅AYG,
together with a pharmaceutically acceptable carrier, excipient or diluent.

73. (New) A method of promoting intestinal function, comprising the step of administering a peptide according to claim 64 to a subject in need of such treatment.

74. (New) A method according to claim 73, in which the subject is suffering from injury to the bowel, an inflammatory condition of the bowel, or short bowel syndrome, has undergone a partial or complete resection of the bowel, or is undergoing total parenteral nutrition.

75. (New) A method of screening of candidate metal ion-binding compounds for ability to modulate the activity of non-amidated gastrins, comprising the steps of

(a) assessing the ability of the compound to inhibit binding of ferric ions to a non-amidated gastrin and/or
(b) assessing the ability of the compound to modulate proliferation and/or migration of cells of a gastric mucosal cell line in response to a non-amidated gastrin.

76. (New) A method according to claim 75, in which the non-amidated gastrin is Ggly₁₇.

**EXPRESS MAIL LABEL NO. EL 979108920 US
ATTORNEY DOCKET NO. 07082.0016U1
Application No. PCT/AU2004/000474**

77. (New) A method according to claim 75, in which the gastric mucosal cell line is IMGE-5.

78. (New) A method according to claim 75, in which the compound is additionally assessed for its ability to inhibit Gamide-induced inositol phosphate production, and/or cellular proliferation in cells which express the CCK-2 receptor.

79. (New) A composition comprising a complex according to claim 69, together with a pharmaceutically acceptable carrier, excipient or diluent.

80. (New) A method of promoting intestinal function, comprising the step of administering

- (a) A peptide which is a fragment of a non-amidated gastrin and which
 - (i) comprises at least glutamate residue 7 of the -(Glu)₅- sequence of non amidated gastrin, and
 - (ii) is capable of binding one or more ferric ions, with the proviso that the peptide is not full length Ggly, full length glycine-extended gastrin or full length progastrin, and
 - (b) a complex comprising
 - (i) a non-amidated gastrin, a peptide fragment thereof according to claim 29, or LE₅AYG, and
 - (ii) a trivalent metal ion
- to a subject in need of such treatment.

81. (New) A method according to claim 73, in which the non-amidated gastrin is Ggly₁₇.